

| L Number | Hits | Search Text | DB | Time stamp |
|----------|------|---------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------|------------------|
| 1 | 9578 | plasminogen near3 (activator or activation) | USPAT; US-PGPUB; EPO; JPO; DERWENT | 2002/12/22 21:53 |
| 7 | 3581 | streptokinase | USPAT; US-PGPUB; EPO; JPO; DERWENT | 2002/12/22 21:54 |
| 13 | 7832 | fibronectin | USPAT; US-PGPUB; EPO; JPO; DERWENT | 2002/12/22 21:54 |
| 19 | 967 | fibrin near4 (bind or binding) | USPAT; US-PGPUB; EPO; JPO; DERWENT | 2002/12/22 21:55 |
| 25 | 119 | ((plasminogen near3 (activator or activation)) and streptokinase and fibronectin and (fibrin near4 (bind or binding))) | USPAT; US-PGPUB; EPO; JPO; DERWENT | 2002/12/22 21:56 |
| 31 | 119 | ((plasminogen near3 (activator or activation)) and streptokinase and fibronectin and (fibrin near4 (bind or binding))) and (fusion protein) | USPAT; US-PGPUB; EPO; JPO; DERWENT | 2002/12/22 21:57 |
| 37 | 0 | streptokinase near5 fibronectin and "129" | USPAT; US-PGPUB; EPO; JPO; DERWENT | 2002/12/22 21:58 |
| 43 | 3 | streptokinase near5 fibronectin and (fibrin near4 (bind or binding)) | USPAT; US-PGPUB; EPO; JPO; DERWENT | 2002/12/22 21:59 |
| 49 | 19 | streptokinase near8 fibronectin | USPAT; US-PGPUB; EPO; JPO; DERWENT | 2002/12/22 21:59 |
| 55 | 2 | ((streptokinase near8 fibronectin) near8 (fibrin near4 (bind or binding))) | USPAT; US-PGPUB; EPO; JPO; DERWENT | 2002/12/22 21:59 |

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=> s streptokinase (8A) fibronectin

41 FILES SEARCHED...

80 FILES SEARCHED...

L1 112 STREPTOKINASE (8A) FIBRONECTIN

=> s fibrin (4A) (bind or binding)

23 FILES SEARCHED...

53 FILES SEARCHED...

90 FILES SEARCHED...

L2 8002 FIBRIN (4A) (BIND OR BINDING)

=> s l1 (8A) l2

58 FILES SEARCHED...

L3 33 L1 (8A) L2

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L5 31 L4 AND STREPTOKINASE

=> d l5 1-31 bib ab

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AN 2000-13074 BIOTECHABS
TI Hybrid **streptokinase**-fibrin binding domain proteins useful for
thrombolytic therapy comprises a streptokinase fused with fibrin binding
domains of human fibronectin having independent fibrin binding domains of
human fibronectin;
vector-mediated gene transfer and expression in host cell
AU Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA CSIR-New-Delhi
LO New Delhi, India.
PI EP 1024192 2 Aug 2000
AI EP 1999-310541 23 Dec 1999
PRAI IN 1998-382598 24 Dec 1998
DT Patent
LA English
OS WPI: 2000-516032 [47]
AB A hybrid plasminogen activator (PA) containing a **streptokinase**
fused with **fibrin binding** domains of human
fibronectin having independent **fibrin binding**
ability and delayed plasminogen activation. The hybrid PA possess the
ability to bind with fibrin independently and also characteristically
retains a PG activation ability which becomes evident only after a
pronounced duration or lag after exposure of the PA to a suitable animal
or human PG Also claimed are: a DNA segment encoding the hybrid PA; an
expression vector; and prokaryotic or eukaryotic cells, transfrome or
transfected with the vector. The hybrid **streptokinase**-fibrin
binding domain proteins are useful in thrombolytic therapy for various
kinds of cardiovascular disorders. (58pp)

L5 ANSWER 2 OF 31 CAPLUS COPYRIGHT 2002 ACS
AN 1992:443664 CAPLUS
DN 117:43664
TI Polypeptides containing the fibrin-binding domain of fibronectin, their
recombinant production, and their use in imaging and therapy
IN Vogel, Tikva; Levanon, Avigdor; Werber, Moshe; Guy, Rachel; Panet, Amos;
Hartman, Jacob; Shaked, Hadassa
PA Bio-Technology General Corp., USA
SO PCT Int. Appl., 192 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 2

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| | AU 660618 | B2 | 19950706 | | |
| | JP 05508766 | T2 | 19931209 | JP 1991-511197 | 19910521 |
| | HU 66189 | A2 | 19941028 | HU 1992-3516 | 19910521 |
| | HU 216302 | B | 19990628 | | |
| | EP 651799 | A1 | 19950510 | EP 1991-911888 | 19910521 |
| | EP 651799 | B1 | 19990818 | | |
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| | AT 183545 | E | 19990915 | AT 1991-911888 | 19910521 |
| | ES 2137928 | T3 | 20000101 | ES 1991-911888 | 19910521 |
| | NO 9204405 | A | 19930113 | NO 1992-4405 | 19921113 |
| | US 5455158 | A | 19951003 | US 1993-58241 | 19930504 |
| | US 5679320 | A | 19971021 | US 1994-259569 | 19940614 |
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| | WO 1991-US3584 | A | 19910521 | | |
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| | US 1994-259569 | A3 | 19940614 | | |
| | US 1995-409750 | A3 | 19950324 | | |

AB Polypeptides having amino acid sequences substantially present in the fibrin-binding domain (FBD) of human fibronectin are labeled with an imageable marker and used in imaging a thrombus or atherosclerotic plaque. Thrombolytic agents bound to the FBD polypeptides are also claimed. Wounds are treated with fusion products of the FBD polypeptide and a polypeptide comprising the cell-binding domain of human fibronectin. A human fibronectin cDNA library was prep'd. and used in cloning and making various FBD polypeptides. The polypeptides were modified with DTPA and radiolabeled with ¹¹¹In and shown to bind to preformed thrombi and thrombi in vivo. They gave a high thrombus:blood ratio of 80-200 after 24 h. The bacterial binding domain of fibronectin was shown to be sep'd. from the FBD since a 31-kDa recombinant FBD polypeptide contg. the entire FBD (residues 1-262 of fibronectin) bound to Staphylococcus aureus, while 18.5 kDa and 12 kDa polypeptides contg. the 1st 154 and 109 amino acid residues of fibronectin, resp., did not. The 18.5 and 12 kDa polypeptides had a high covalent binding specificity for fibrin together with a narrower spectrum of activities and lower specificity for other ligands such as vascular components and bacteria than the 31 kDa protein which is advantageous for thrombus imaging.

L5 ANSWER 3 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAY90282 Protein DGENE

TI Hybrid **streptokinase-fibrin binding** domain
polypeptides useful for thrombolytic therapy comprises a
streptokinase fused with **fibrin binding**
domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802

58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents the human *Streptococcus equisimilis* **streptokinase** protein sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 4 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAY90281 Protein DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a human fibronectin fragment, containing fibrin binding domains. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use

of streptokinase.

L5 ANSWER 5 OF 31 DGENE (C) 2002 THOMSON DERWENT
AN AAY90280 Peptide DGENE
TI Hybrid **streptokinase-fibrin binding** domain
polypeptides useful for thrombolytic therapy comprises a
streptokinase fused with **fibrin binding**
domains of human **fibronectin** -
IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.
PI EP 1024192 A2 20000802 58p
AI EP 1999-310541 19991223
PRAI IN 1998-3825 19981224
DT Patent
LA English
OS 2000-516032 [47]
AB This sequence represents the intergenic region of a chimeric
streptokinase-fibrin binding domain (SK-FBD) protein sequence. The
invention relates to a hybrid plasminogen activator (PA) comprises a
polypeptide fusion between **streptokinase** (SK), which are
capable of plasminogen (PG) activation, and fibrin binding regions of
human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or
1 and 2. The hybrid PA possesses the ability to bind with fibrin
independently and also characteristically retains a PG activation ability
which becomes evident only after a pronounced duration, or lag, after
exposure of the PA to a suitable animal or human PG. The hybrid
streptokinase-fibrin binding domain polypeptides are useful in
thrombolytic therapy for various kinds of cardiovascular disorders. The
hybrids have enhanced fibrin selectivity as well as kinetics of
plasminogen activation that are distinct from that of natural
streptokinase in being characterised by a temporary delay, or lag
of several minutes in the natural rate of the catalytic conversion of
plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins
can bind tightly with fibrin in blood clots soon after introduction into
the vascular system without significantly activating the circulating
blood plasminogen to plasmin, thus aiding in the localisation of the
plasminogen activation process to the site of pathological thrombus. This
overcomes systemic plasminogen activation encountered during clinical use
of **streptokinase**.

L5 ANSWER 6 OF 31 DGENE (C) 2002 THOMSON DERWENT
AN AAA37644 DNA DGENE
TI Hybrid **streptokinase-fibrin binding** domain
polypeptides useful for thrombolytic therapy comprises a
streptokinase fused with **fibrin binding**
domains of human **fibronectin** -
IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.
PI EP 1024192 A2 20000802 58p
AI EP 1999-310541 19991223
PRAI IN 1998-3825 19981224
DT Patent
LA English
OS 2000-516032 [47]
AB This sequence represents a chimeric streptokinase-fibrin binding domain
(SK-FBD) protein coding sequence. The invention relates to a hybrid
plasminogen activator (PA) comprises a polypeptide fusion between
streptokinase (SK), which are capable of plasminogen (PG)
activation, and fibrin binding regions of human fibronectin, which are
from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA
possesses the ability to bind with fibrin independently and also
characteristically retains a PG activation ability which becomes evident

only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 7 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37643 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a chimeric streptokinase-fibrin binding domain (SK-FBD) protein coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 8 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37642 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a chimeric streptokinase-fibrin binding domain (SK-FBD) protein coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 9 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37641 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence is a PCR primer used in the construction of a chimeric streptokinase-fibrin binding domain (SK-FBD) protein coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This

overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 10 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37640 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain
polypeptides useful for thrombolytic therapy comprises a
streptokinase fused with **fibrin binding**
domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M

PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence is a PCR primer used in the construction of a chimeric streptokinase-fibrin binding domain (SK-FBD) protein coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase-fibrin binding domain** polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 11 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37639 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain
polypeptides useful for thrombolytic therapy comprises a
streptokinase fused with **fibrin binding**
domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M

PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence is a PCR primer used in the construction of a chimeric streptokinase-fibrin binding domain (SK-FBD) protein coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin

independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 12 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37638 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence is a PCR primer used in the construction of a chimeric streptokinase-fibrin binding domain (SK-FBD) protein coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 13 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37637 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M

PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802

58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a chimeric streptokinase-fibrin binding domain (SK-FBD) protein coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 14 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37636 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M

PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802

58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a PCR primer for the Streptococcus equisimilis **streptokinase** (SK) coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without

significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 15 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37635 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain
polypeptides useful for thrombolytic therapy comprises a
streptokinase fused with **fibrin binding**
domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a PCR primer for the *Streptococcus equisimilis* **streptokinase** (SK) coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase-fibrin** binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 16 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37634 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain
polypeptides useful for thrombolytic therapy comprises a
streptokinase fused with **fibrin binding**
domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a PCR primer for the *Streptococcus equisimilis* **streptokinase** (SK) coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA

possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 17 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37633 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192. A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents the human Streptococcus equisimilis **streptokinase** coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 18 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37632 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p
AI EP 1999-310541 19991223
PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a human fibronectin coding sequence fragment, containing fibrin binding domains. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 19 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37631 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a PCR primer for the human fibrin binding domain coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the

plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 20 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37630 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a PCR primer for the human fibrin binding domain coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase-fibrin binding** domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 21 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37629 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents the intergenic region of a chimeric streptokinase-fibrin binding domain (SK-FBD) protein coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin

independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 22 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37628 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a **streptokinase**-NTR (SK-NTR) gene (where NTR stands for N-terminally repaired with native sequence). The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 23 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37627 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M

PA (COUL) CSIR COUNCIL SCI IND RES.
PI EP 1024192 A2 20000802 58p
AI EP 1999-310541 19991223
PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a PCR primer for the *Streptococcus equisimilis* **streptokinase** (SK) coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 24 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37626 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a fragment of the *Streptococcus equisimilis* **streptokinase** (SK) coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without

significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 25 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37625 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain
polypeptides useful for thrombolytic therapy comprises a
streptokinase fused with **fibrin binding**
domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a fragment of the *Streptococcus equisimilis* **streptokinase** (SK) coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase-fibrin** binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 26 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37624 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain
polypeptides useful for thrombolytic therapy comprises a
streptokinase fused with **fibrin binding**
domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a PCR primer for the *Streptococcus equisimilis* **streptokinase** (SK) coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA

possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 27 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37623 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a PCR primer for the Streptococcus equisimilis **streptokinase** (SK) coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 28 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37622 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a **streptokinase**-NTRN (SK-NTRN) gene (where NTRN stands for N-terminally repaired with native sequence). The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 29 OF 31 DGENE (C) 2002 THOMSON DERWENT

AN AAA37621 DNA DGENE

TI Hybrid **streptokinase-fibrin binding** domain polypeptides useful for thrombolytic therapy comprises a **streptokinase** fused with **fibrin binding** domains of human **fibronectin** -

IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.

PI EP 1024192 A2 20000802 58p

AI EP 1999-310541 19991223

PRAI IN 1998-3825 19981224

DT Patent

LA English

OS 2000-516032 [47]

AB This sequence represents a PCR primer for the *Streptococcus equisimilis* **streptokinase** (SK) coding sequence. The invention relates to a hybrid plasminogen activator (PA) comprises a polypeptide fusion between **streptokinase** (SK), which are capable of plasminogen (PG) activation, and fibrin binding regions of human fibronectin, which are from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA possesses the ability to bind with fibrin independently and also characteristically retains a PG activation ability which becomes evident only after a pronounced duration, or lag, after exposure of the PA to a suitable animal or human PG. The hybrid **streptokinase**-fibrin binding domain polypeptides are useful in thrombolytic therapy for various kinds of cardiovascular disorders. The hybrids have enhanced fibrin selectivity as well as kinetics of plasminogen activation that are distinct from that of natural **streptokinase** in being characterised by a temporary delay, or lag of several minutes in the natural rate of the catalytic conversion of plasminogen to plasmin (i.e. delayed-action thrombolysis). The proteins can bind tightly with fibrin in blood clots soon after introduction into the vascular system without

significantly activating the circulating blood plasminogen to plasmin, thus aiding in the localisation of the plasminogen activation process to the site of pathological thrombus. This overcomes systemic plasminogen activation encountered during clinical use of **streptokinase**.

L5 ANSWER 30 OF 31 DGENE (C) 2002 THOMSON DERWENT
AN AAA37620 DNA DGENE
TI Hybrid **streptokinase-fibrin binding** domain
polypeptides useful for thrombolytic therapy comprises a
streptokinase fused with **fibrin binding**
domains of human **fibronectin** -
IN Sahni G; Kumar R; Roy C; Rajogopal K; Nihalani D; Sundaram V; Yadav M
PA (COUL) CSIR COUNCIL SCI IND RES.
PI EP 1024192 A2 20000802 58p
AI EP 1999-310541 19991223
PRAI IN 1998-3825 19981224
DT Patent
LA English
OS 2000-516032 [47]
AB This sequence represents a PCR primer for the Streptococcus equisimilis
streptokinase (SK) coding sequence. The invention relates to a
hybrid plasminogen activator (PA) comprises a polypeptide fusion between
streptokinase (SK), which are capable of plasminogen (PG)
activation, and fibrin binding regions of human fibronectin, which are
from fibrin binding domains (FBD) 4 and 5 or 1 and 2. The hybrid PA
possesses the ability to bind with fibrin independently and also
characteristically retains a PG activation ability which becomes evident
only after a pronounced duration, or lag, after exposure of the PA to a
suitable animal or human PG. The hybrid **streptokinase-fibrin**
binding domain polypeptides are useful in thrombolytic therapy for
various kinds of cardiovascular disorders. The hybrids have enhanced
fibrin selectivity as well as kinetics of plasminogen activation that are
distinct from that of natural **streptokinase** in being
characterised by a temporary delay, or lag of several minutes in the
natural rate of the catalytic conversion of plasminogen to plasmin (i.e.
delayed-action thrombolysis). The proteins can bind tightly with fibrin
in blood clots soon after introduction into the vascular system without
significantly activating the circulating blood plasminogen to plasmin,
thus aiding in the localisation of the plasminogen activation process to
the site of pathological thrombus. This overcomes systemic plasminogen
activation encountered during clinical use of **streptokinase**.

L5 ANSWER 31 OF 31 USPATFULL
AN 92:80812 USPATFULL
TI Pharmaceutically active conjugates having improved body tissue binding
specificity
IN Brown, Robert A., St. Albans, Great Britain
PA Central Blood Laboratories Authority, Borehamwood, Great Britain
(non-U.S. corporation)
PI US 5151412 19920929
WO 8803810 19880602
AI US 1989-359662 19890721 (7)
WO 1987-GB854 19871127
19890721 PCT 371 date
19890721 PCT 102(e) date
PRAI GB 1986-28398 19861127
DT Utility
FS Granted
EXNAM Primary Examiner: Nucker, Christine M.; Assistant Examiner: Kim, Kay K.
LREP Foley & Lardner
CLMN Number of Claims: 18
ECL Exemplary Claim: 1

DRWN 3 Drawing Figure(s); 3 Drawing Page(s)

LN.CNT 1358

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Pharmaceutically active conjugates comprising a pharmaceutically active substance for treating a disorder of the body that involves a specified body tissue conjugated directly or indirectly with at least one fragment of an adhesive glycoprotein such as fibronectin, the said glycoprotein fragment(s) having improved binding specificity compared with the parent protein for the said body tissue.